

REMARKS

Entry of the foregoing amendment is respectfully requested.

Claims 14-16, added in the Second Preliminary Amendment on October 1, 2001, were incorrectly numbered. The second claim 14 has been renumbered at claim 15, and the remaining number of claims have been corrected.

In response to the Election Requirement, applicants hereby elect Species A, directed to compounds of formula (1) with A as a monocyclic aryl. This election is made for examination purposes only.

Claims readable on the elected species include claims 1-18.

In view of the above, it is respectfully requested that the restriction requirement be withdrawn or at the very least altered.

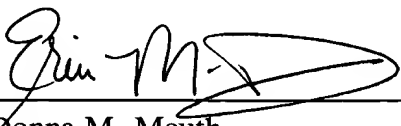
In the event that there are any questions relating to this amendment or the application in general, it would be appreciated if the Examiner would contact the undersigned attorney at (508) 339-3684.

Early and favorable action in the form of a notice of allowance is respectfully requested.

Respectfully submitted,

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Date: January 23, 2003

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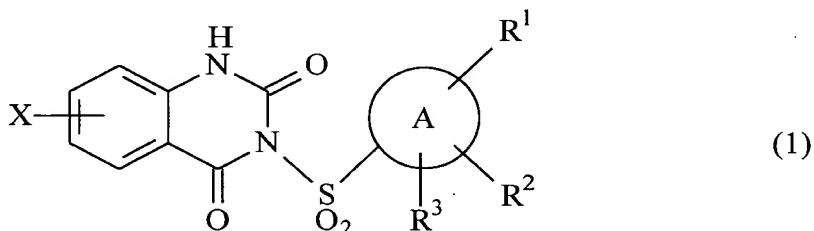


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Marked-up Claims 14-17

[14] 15. (Amended) A method for suppressing lipid deposition in a blood vessel comprising administering to a patient in need of such treatment a chymase inhibitor in an amount effective for suppressing lipid deposition in the blood vessel.

[15] 16. (Amended) A method according to claim [14] 15, wherein the chymase inhibitor is a quinazoline derivative represented by the formula (1):



wherein, the ring A represents an aryl ring,

R¹ represents a hydroxy group, an amino group, or a lower alkylamino group having 1 to 4 carbons that may be substituted with a carboxylic group, a lower aralkylamino group having 7 to 10 carbons that may be substituted with a carboxylic group, an amino group acylated with a lower fatty acid having 1 to 4 carbons that may be substituted with a carboxylic group, an amino group acylated with an aromatic carboxylic acid that may be substituted with a carboxylic group, an amino group acylated with a

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heteroaromatic carboxylic acid that may be substituted with a carboxylic group, an amino group sulfonylated with a lower alkanesulfonic acid having 1 to 4 carbons that may be substituted with a carboxylic group, an amino group sulfonylated with an aromatic sulfonic acid that may be substituted with a carboxylic group, an amino group sulfonylated with a heteroaromatic sulfonic acid that may be substituted with a carboxylic group, a lower alkyl group having 1 to 4 carbons substituted with a carboxylic group, or a lower alkylene group having 2 to 4 carbons substituted with a carboxylic group;

R^2 and R^3 , which may be the same or different, represent a hydrogen, a lower alkyl group having 1 to 4 carbons that may be substituted, a halogen atom, a hydroxy group, a lower alkoxy group having 1 to 4 carbons, an amino group, a lower alkylamino group having 1 to 4 carbons that may be substituted, a lower aralkylamino group having 7 to 10 carbons that may be substituted, an amino group acylated with a lower fatty acid having 1 to 4 carbons that may be substituted with a carboxylic group, an amino group acylated with an aromatic carboxylic acid that may be substituted with a carboxylic group, an amino group acylated with a heteroaromatic carboxylic acid that may be substituted with a carboxylic group, an amino group sulfonylated with a lower alkanesulfonic acid having 1 to 4 carbons that may be substituted with a carboxylic group, an amino group sulfonylated with an aromatic sulfonic acid that may be substituted with a carboxylic group, an amino group sulfonylated with a heteroaromatic sulfonic acid that may be substituted with a carboxylic group, or a carboxylic group; or

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when the ring A is a benzene ring, R¹ and R², together with the benzene ring to be substituted, may form a fused heterocyclic ring that may be substituted with a carboxylic acid, and a carbon atom in said fused heterocyclic ring may form a carbonyl group wherein R³ is as defined above; and

X represents a hydrogen atom, a lower alkyl group having 1-4 carbons, a lower alkoxy group having 1 to 4 carbons, a halogen atom, a hydroxy group, an amino group, or a nitro group;

or a pharmaceutically acceptable salt thereof.

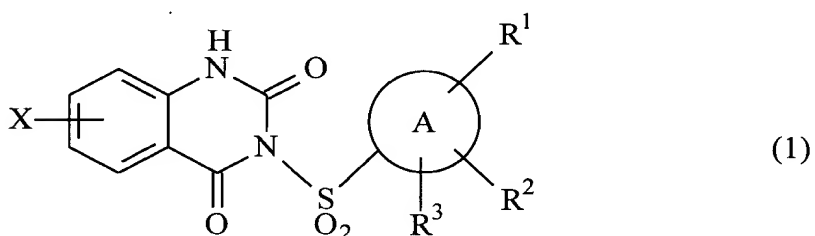
[16] 17. (Amended) A method for preventing or treating a disease accompanied by abnormal vascular function in which lipid deposition in the blood vessel is involved,

wherein said disease is selected from the group consisting of arteriosclerosis, cardiac acute coronary syndrome, restenosis after percutaneous transluminal coronary angioplasty, obstructive arteriosclerosis, obstructive thrombotic vasculitis, atherosclerosis, cerebral infarction, intermittent claudication, lower limb gangrene, renal vascular hypertension, renal arterial aneurysm and renal infarction,

comprising administering to a patient in need of such treatment a quinazoline derivative represented by the formula (1):

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wherein, the ring A represents an aryl ring,

R¹ represents a hydroxy group, an amino group, or a lower alkylamino group having 1 to 4 carbons that may be substituted with a carboxylic group, a lower aralkylamino group having 7 to 10 carbons that may be substituted with a carboxylic group, an amino group acylated with a lower fatty acid having 1 to 4 carbons that may be substituted with a carboxylic group, an amino group acylated with an aromatic carboxylic acid that may be substituted with a carboxylic group, an amino group acylated with a heteroaromatic carboxylic acid that may be substituted with a carboxylic group, an amino group sulfonylated with a lower alkanesulfonic acid having 1 to 4 carbons that may be substituted with a carboxylic group, an amino group sulfonylated with an aromatic sulfonic acid that may be substituted with a carboxylic group, an amino group sulfonylated with a heteroaromatic sulfonic acid that may be substituted with a carboxylic group, a lower alkyl group having 1 to 4 carbons substituted with a carboxylic group, or a lower alkylene group having 2 to 4 carbons substituted with a carboxylic group;

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R^2 and R^3 , which may be the same or different, represent a hydrogen, a lower alkyl group having 1 to 4 carbons that may be substituted, a halogen atom, a hydroxy group, a lower alkoxy group having 1 to 4 carbons, an amino group, a lower alkylamino group having 1 to 4 carbons that may be substituted, a lower aralkylamino group having 7 to 10 carbons that may be substituted, an amino group acylated with a lower fatty acid having 1 to 4 carbons that may be substituted with a carboxylic group, an amino group acylated with an aromatic carboxylic acid that may be substituted with a carboxylic group, an amino group acylated with a heteroaromatic carboxylic acid that may be substituted with a carboxylic group, an amino group sulfonylated with a lower alkanesulfonic acid having 1 to 4 carbons that may be substituted with a carboxylic group, an amino group sulfonylated with an aromatic sulfonic acid that may be substituted with a carboxylic group, an amino group sulfonylated with a heteroaromatic sulfonic acid that may be substituted with a carboxylic group, or a carboxylic group; or

when the ring A is a benzene ring, R^1 and R^2 , together with the benzene ring to be substituted, may form a fused heterocyclic ring that may be substituted with a carboxylic acid, and a carbon atom in said fused heterocyclic ring may form a carbonyl group wherein R^3 is as defined above; and

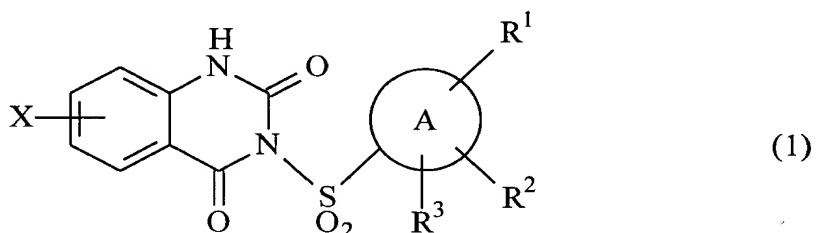
X represents a hydrogen atom, a lower alkyl group having 1-4 carbons, a lower alkoxy group having 1 to 4 carbons, a halogen atom, a hydroxy group, an amino group, or a nitro group;

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or a pharmaceutically acceptable salt thereof,
in an amount effective for treating said diseases accompanied by abnormal vascular function.

[17] 18. (Amended) A method for suppressing lipid deposition in a blood vessel comprising administering to a patient in need of such treatment a quinazoline derivative represented by the formula (1):



wherein, the ring A represents an aryl ring,

R¹ represents a hydroxy group, an amino group, or a lower alkylamino group having 1 to 4 carbons that may be substituted with a carboxylic group, a lower aralkylamino group having 7 to 10 carbons that may be substituted with a carboxylic group, an amino group acylated with a lower fatty acid having 1 to 4 carbons that may be substituted with a carboxylic group, an amino group acylated with an aromatic carboxylic acid that may be substituted with a carboxylic group, an amino group acylated with a

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heteroaromatic carboxylic acid that may be substituted with a carboxylic group, an amino group sulfonylated with a lower alkanesulfonic acid having 1 to 4 carbons that may be substituted with a carboxylic group, an amino group sulfonylated with an aromatic sulfonic acid that may be substituted with a carboxylic group, an amino group sulfonylated with a heteroaromatic sulfonic acid that may be substituted with a carboxylic group, a lower alkyl group having 1 to 4 carbons substituted with a carboxylic group, or a lower alkylene group having 2 to 4 carbons substituted with a carboxylic group;

R^2 and R^3 , which may be the same or different, represent a hydrogen, a lower alkyl group having 1 to 4 carbons that may be substituted, a halogen atom, a hydroxy group, a lower alkoxy group having 1 to 4 carbons, an amino group, a lower alkylamino group having 1 to 4 carbons that may be substituted, a lower aralkylamino group having 7 to 10 carbons that may be substituted, an amino group acylated with a lower fatty acid having 1 to 4 carbons that may be substituted with a carboxylic group, an amino group acylated with an aromatic carboxylic acid that may be substituted with a carboxylic group, an amino group acylated with a heteroaromatic carboxylic acid that may be substituted with a carboxylic group, an amino group sulfonylated with a lower alkanesulfonic acid having 1 to 4 carbons that may be substituted with a carboxylic group, an amino group sulfonylated with an aromatic sulfonic acid that may be substituted with a carboxylic group, an amino group sulfonylated with a heteroaromatic sulfonic acid that may be substituted with a carboxylic group, or a carboxylic group; or

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when the ring A is a benzene ring, R¹ and R², together with the benzene ring to be substituted, may form a fused heterocyclic ring that may be substituted with a carboxylic acid, and a carbon atom in said fused heterocyclic ring may form a carbonyl group wherein R³ is as defined above; and

X represents a hydrogen atom, a lower alkyl group having 1-4 carbons, a lower alkoxy group having 1 to 4 carbons, a halogen atom, a hydroxy group, an amino group; or a nitro group;

or a pharmaceutically acceptable salt thereof,

in an amount effective for suppressing lipid deposition in the blood vessel.